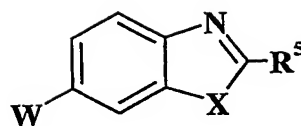


WE CLAIM:

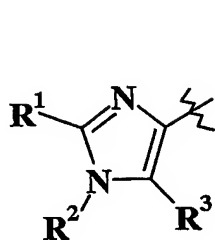
1. A compound of Formula I:



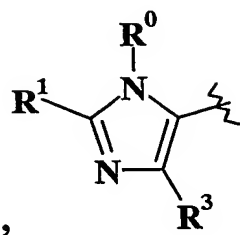
I

where:

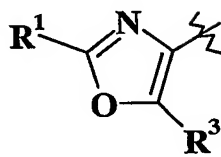
W is



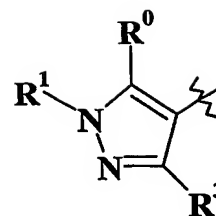
(i)



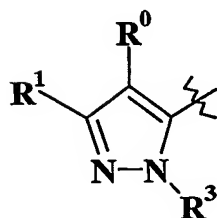
(ii)



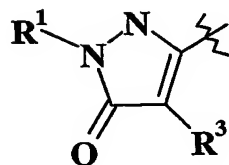
(iii)



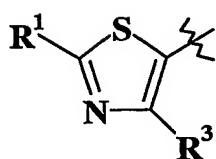
(iv)



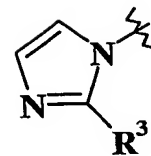
(v)



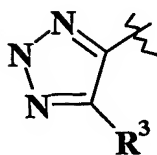
(vi)



(vii)

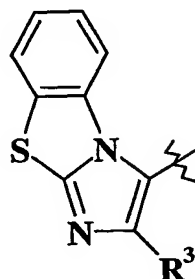


(viii)



(ix)

, or



(x)

;

$R^0$  is

(a) selected from the group consisting of hydrogen,  $C_1$ - $C_6$  alkyl, cyano, ( $C_1$ - $C_4$  alkylene)- $R^{11}$ , 3-hydroxyprop-2-yl, (1-phenyl)-2-hydroxyeth-1-yl, (1-cyclohexyl)-3-hydroxyprop-2-yl, 4-methoxybenzyl, 1,4-dioxoaspiro[4,5]dec-8-yl, tetrahydropyran, 2,2,6,6-tetramethylpiperidin-4-yl, and cyclohexan-1-on-4-yl,

(b) phenyl optionally substituted with one substituent selected from the group consisting of nitro and amino,

(c) piperidin-4-yl optionally substituted with one substituent selected from the group consisting of  $C_1$ - $C_4$  alkyl,  $C_1$ - $C_4$  alkoxy carbonyl, and benzyl, or

(d)  $C_3$ - $C_6$  cycloalkyl optionally substituted with one substituent selected from the group consisting of  $C_1$ - $C_4$  alkoxy carbonylamino, amino, hydroxy, and  $C_1$ - $C_4$  alkylene-OH;

$R^1$  is

(a) selected from the group consisting of hydrogen,  $C_1$ - $C_6$  alkyl,  $C_2$ - $C_4$  alkynyl, halo, amino, azido, formyl, 1-( $C_1$ - $C_4$  alkoxy carbonyl)ethen-2-yl, 1-( $C_1$ - $C_4$  alkoxy carbonyl)ethyl, 1-( $C_1$ - $C_4$  carboxy)ethyl, ( $C_1$ - $C_4$  alkylene)benzyloxy, trifluoromethyl, trimethylsilylethynyl, but-3-yn-1-ol,  $C_3$ - $C_6$  cycloalkyl, tetrahydropyran-4-yl, hydroxymethyl, 2-(piperidin-1-yl)methyl,  $N,N',N'$ -[trimethyl]-2-(aminoethylamino)methyl, (morpholin-4-yl)methyl, dimethylaminomethyl,  $N$ -[2-(piperidin-1-yl)eth-1-yl]-aminomethyl,  $N',N'$ -dimethyl-2-(aminoethylamino)methyl, pyridinyl, thiazolyl, triazolyl, benzo(1,3)dioxolan-5-yl, and imidazol-2-yl,

(b) phenyl optionally substituted with one to three substituents independently selected from the group consisting of  $C_1$ - $C_4$  alkyl, halo, nitro, amino,  $C_1$ - $C_4$  alkoxy, trifluoromethyl, trifluoromethoxy, trifluoromethylsulfanyl, methylsulfonyl, methylsulfonamidyl, pyrrolidin-1-yl, morpholin-4-yl, 4-( $C_1$ - $C_4$  alkyl)piperazin-1-yl,  $-NR^6R^7$ , and

$C_1$ - $C_4$  alkoxy optionally substituted with one substituent selected from the group consisting of piperidin-1-yl, pyrrolidin-1-yl, morpholin-4-yl, azepin-4-yl, and di( $C_1$ - $C_4$  alkyl)amino,

(c) thienyl optionally substituted with one substituent selected from the group consisting of halo, nitro, amino, and C<sub>1</sub>-C<sub>4</sub> alkyl, or

(d) piperidin-4-yl optionally substituted at the 1-position from the group consisting of C<sub>1</sub>-C<sub>4</sub> alkyl, C<sub>1</sub>-C<sub>4</sub> alkoxy carbonyl, benzyloxycarbonyl, and (C<sub>1</sub>-C<sub>4</sub> alkylene)-R<sup>8</sup>;

Alternatively R<sup>0</sup> and R<sup>1</sup> may be taken together to form a fully saturated C<sub>3</sub>-C<sub>4</sub> carbon chain or a fully unsaturated C<sub>3</sub>-C<sub>4</sub> carbon chain optionally substituted with halo or C<sub>1</sub>-C<sub>4</sub> alkyl;

**R<sup>2</sup> is hydrogen, C<sub>1</sub>-C<sub>4</sub> alkyl, or benzyl;**

R<sup>3</sup> is thienyl or phenyl optionally substituted with one to two substituents independently selected from the group consisting of halo, C<sub>1</sub>-C<sub>4</sub> alkyl, C<sub>1</sub>-C<sub>4</sub> alkoxy, and trifluoromethyl;

R<sup>4</sup> is hydrogen, (C<sub>1</sub>-C<sub>4</sub> alkyl)sulfonyl, or (C<sub>3</sub>-C<sub>6</sub> cycloalkyl)sulfonyl; or (C<sub>1</sub>-C<sub>4</sub> alkyl)<sub>2</sub>N-sulfonyl;

$R^5$  is halo, hydrogen, or  $-NR^9R^{10}$ ;

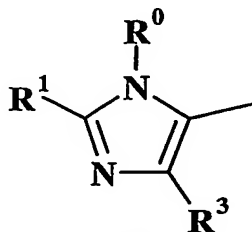
R<sup>6</sup> and R<sup>7</sup> are individually at each occurrence selected from hydrogen, carbonyl, or C<sub>1</sub>-C<sub>4</sub> alkyl providing that at least one of R<sup>6</sup> and R<sup>7</sup> is hydrogen;

R<sup>8</sup> is hydroxy, trifluoromethyl, dimethylamino, phenyl, pyridinyl, or 1-methylimidazol-2-yl,;

**R<sup>9</sup> is independently at each instance hydrogen or C<sub>1</sub>-C<sub>4</sub> alkyl;**

R<sup>10</sup> is hydrogen, C<sub>1</sub>-C<sub>4</sub> alkyl, or benzyl;

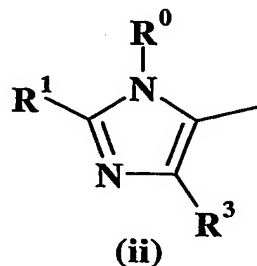
R<sup>11</sup> is C<sub>1</sub>-C<sub>4</sub> alkoxy, hydroxy, C<sub>1</sub>-C<sub>4</sub> alkoxycarbonyl, C<sub>1</sub>-C<sub>4</sub> alkoxycarbonylamino, C<sub>3</sub>-C<sub>6</sub> cycloalkyl, phenyl optionally substituted with one to two substituents independently selected from the group consisting of C<sub>1</sub>-C<sub>4</sub> alkoxy and halo, morpholin-4-yl, or pyridinyl;



provided that when  $W$  is (ii) then

(a) at least one of  $R^0$  and  $R^1$  is hydrogen or  $C_1$ - $C_6$  alkyl; or

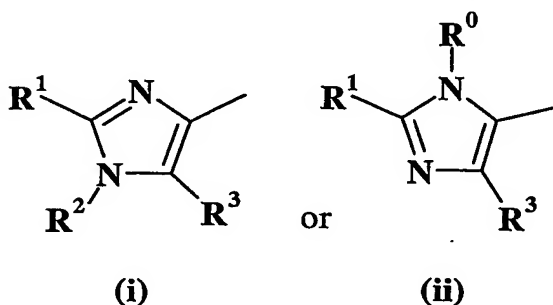
- (b)  $R^0$  and  $R^1$  may be taken together to form a fully saturated  $C_3-C_4$  carbon chain or a fully unsaturated  $C_3-C_4$  carbon chain optionally substituted with halo or  $C_1-C_4$  alkyl;



also provided that when X is S, W is

- 5 or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

2. A compound of Claim 1, W is is either



- 10 3. A compound of either of Claims 1 or 2, where X is N(isopropylsulfonyl) and  $R^5$  is  $-NH_2$ .

4. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(2,6-difluorophenyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically  
15 acceptable salt or a pharmaceutically acceptable solvate thereof.

5. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(4-chlorophenyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

6. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(2-chloro-6-fluorophenyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

5 7. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(*tert*-butyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

10 8. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(2-chloro-6-fluorophenyl)-5-(4-fluorophenyl))-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

15 9. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(isopropyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

20 10. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(cyclopropyl)-5-(4-fluorophenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

11. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(2-fluoro-6-trifluoromethylphenyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

25 12. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(2-trifluoromethylphenyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

30 13. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(cyclohexyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

14. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(1-(methyl)-2-(2,6-difluorophenyl)-4-(phenyl)-imidazol-5-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

5 15. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(2,6-dichlorophenyl)-5-(4-fluorophenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

10 16. A compound of Claim 1, which is 1-isopropylsulfonyl-2-amino-6-(2-(2,6-dichlorophenyl)-5-(phenyl)-imidazol-4-yl)-benzimidazole or a pharmaceutically acceptable salt or a pharmaceutically acceptable solvate thereof.

15 17. A pharmaceutical formulation comprising a compound of Claim 1 and a pharmaceutically acceptable carrier, diluent, or excipient.

18. A method of inhibiting p-38 kinase in a mammal comprising administering to a mammal in need of such treatment an effective amount of a compound of Claim 1.

20 19. A method of treating conditions resulting from excessive cytokine production in a mammal comprising administering to a mammal in need of such treatment a cytokine-suppressing amount of a compound of Claim 1.

20. A method of Claim 7, where the cytokine is tumor necrosis factor  $\alpha$ .

25 21. A method of inhibiting the growth of a susceptible neoplasm in a mammal comprising administering to a mammal in need of such treatment a p38 inhibiting amount of a compound of Claim 1.

30 22. A method of inhibiting metastasis in a mammal comprising administering to a mammal in need of such treatment a p38 inhibiting amount of a compound of Claim 1.

23. A method of treating rheumatoid arthritis in a mammal comprising administering to a mammal in need of such treatment a p38 inhibiting amount of a compound of Claim 1.